This listing of claims will replace all prior versions, and listings, of claims in the application:

Listing of Claims:

1.   **(Currently Amended)** A method for achieving high sensitivity detection and/or high accuracy quantitation of a plurality of target proteins in a biological sample, the method comprising the steps of:

   (1)   fragmenting proteins in the sample using a predetermined denaturation and proteolytic protocol to generate a solution of polypeptide analytes comprising recognition sequences comprising peptide epitope tags (PETs) displaying solvent accessible binding surfaces unambiguously indicative of the presence in the sample of the proteins from which they are derived;

   (2)   providing an addressable array of plural capture agents, two or more different capture agents for detecting a target protein in a test sample, which capture agents are provided as an addressable array, and each of which capture agents selectively interacts interact with a peptide epitope tag (PET) of said target protein to unambiguouly indicate the presence of said target protein in said sample;

   ([[2]]3)contacting said array with [[a]] said solution of polypeptide analytes to capture polypeptide analytes by binding interaction between respective capture agents and said PETs presented by respective said analytes; and, produced by denaturation and/or cleavage of proteins from the test sample;

   ([[3]]4)detecting the presence and amounts of said target proteins in the sample from the interaction of by detecting said captured polypeptide analytes with each said capture agents secondary capture agents specific for captured polypeptide analytes and labeled with a detectable moiety;

   (4)   quantitating, if present, the amount of the target protein in the sample by averaging the results obtained from each said capture agents in (3).

2.   **(Currently Amended)** The method of claim 1, comprising providing an addressable array of capture agents comprising plural capture agents which selectively interact with different PETs from the same target protein and quantitating, if present, the amount of the target protein in the sample by averaging the results obtained from each said capture

agent ~~wherein each said different capture agents specifically bind a different PET of said target protein~~.

3.   (**Currently Amended**) The method of claim [[2]] 1, wherein said ~~different~~ capture agents <u>comprise antibodies</u> ~~belong to the same category of capture agent~~.

4.   (**Currently Amended**) The method of claim [[3]] 1, wherein said ~~category of~~ capture agent <u>comprise a member selected from the group consisting of</u> ~~includes: antibody,~~ non-antibody polypeptide, PNA (peptide nucleic acids), scaffolded peptide, peptidomimetic compound, polynucleotide, carbohydrates, artificial polymers, plastibody, chimeric binding <u>agent</u> ~~agnet~~ derived from low-affinity ligand, and small organic molecules.

5.   (**Currently Amended**) The method of claim [[2]] 1, wherein <u>said secondary capture agents bind to an epitope on a said polypeptide analyte separate from said solvent accessible binding surfaces</u> ~~at least two of said different capture agents belong to different categorys of capture agent selected from antibody, non-antibody polypeptide, PNA (peptide nucleic acids), scaffolded peptide, peptidomimetic compound, polynucleotide, carbohydrates, artificial polymers, plastibody, chimeric binding agnet derived from low-affinity ligand, and small organic molecules~~.

6.   (**Currently Amended**) The method of claim 1, wherein a subset of said capture agents bind to the same PET~~, and wherein each capture agents of said subset belong to different category of capture agent selected from: antibody, non-antibody polypeptide, PNA (peptide nucleic acids), scaffolded peptide, peptidomimetic compound, polynucleotide, carbohydrates, artificial polymers, plastibody, chimeric binding agnet derived from low-affinity ligand, and small organic molecules~~.

7.   (**Original**) The method of claim 1, wherein said target protein has two or more different forms within said biological sample.

8.   (**Original**) The method of claim 7, wherein said different forms include unprocessed / pro-form and processed / mature form.

9.   (**Original**) The method of claim 7, wherein said different forms include different alternative splicing forms.

10-12. (**Canceled**)

13. **(Currently Amended)** The method of claim [[12]] 7, further comprising determining the percentage of one form of said target protein as compared to the total target protein, or ratio of a first form of said target protein to a second form of said target protein.

14-15. **(Canceled)**

16. **(Original)** The method of claim 1, wherein, for each capture agent, the method has a regression coefficient (R^2) of 0.95 or greater.

17-18. **(Canceled)**

19. **(Original)** The method of claim 1, wherein said sample is a body fluid selected from: saliva, mucous, sweat, whole blood, serum, urine, amniotic fluid, genital fluid, fecal material, marrow, plasma, spinal fluid, pericardial fluid, gastric fluid, abdominal fluid, peritoneal fluid, pleural fluid, synovial fluid, cyst fluid, cerebrospinal fluid, lung lavage fluid, lymphatic fluid, tears, prostatitc fluid, extraction from other body parts, or secretion from other glands; or from supernatant, whole cell lysate, or cell fraction obtained by lysis and fractionation of cellular material, extract or fraction of cells obtained directly from a biological entity or cells grown in an artificial environment.

20. **(Currently Amended)** The method of claim 1, wherein said sample is obtained from human, mouse, rat, frog ~~(Xenopus)~~, fish ~~(zebra-fish)~~, fly ~~(Drosophila melanogaster)~~, nematode ~~(C. elegans)~~, fission or budding yeast, or plant ~~(Arabidopsis thaliana)~~.

21. **(Original)** The method of claim 1, wherein said sample is produced by treatment of membrane bound proteins.

22. **(Canceled)**

23. **(Currently Amended)** The method of claim 1, wherein ~~step (3) is effectuated by using secondary capture agents specific for captured polypeptide analytes, wherein~~ said secondary capture agent is labeled <u>with</u> ~~by~~ a detectable moiety selected from: an enzyme, a fluorescent label, a stainable dye, a chemilumninescent compound, a colloidal particle, a radioactive isotope, a near-infrared dye, a DNA dendrimer, a water-soluble quantum dot, a latex bead, a selenium particle, or a europium nanoparticle.

24. **(Currently Amended)** The method of claim 23, wherein said secondary capture agent is <u>labeled with a fluorophore</u> ~~specific for a post-translational modification~~.

25.   (Canceled)

26.   (Original) The method of claim 1, wherein said sample contains billion molar excess of unrelated proteins or fragments thereof relative to said target protein.

27.   (Original) The method of claim 1, wherein said PET is identified based on one or more of the protein sources selected from: sequenced genome or virtually translated proteome, virtually translated transcriptome, or mass spectrometry database of tryptic fragments.

28.   (Currently Amended) The method of claim 1, wherein one or a combination of said the target proteins serve as [[is]] a biomarker with a concentration of about 1-5 pM in said sample.

29.   (Canceled)

30.   (Currently Amended) An array of capture agents for detecting and quantitating a target protein plural target splice variant proteins within a biological sample, the array comprising a plurality of capture agents, each immobilized on a distinct addressable location on a solid support, plural each of said capture agents specifically binds binding to recognition sequences comprising peptide epitope tags (PETs) displaying solvent accessible binding surfaces unambiguously indicative of the presence in the sample of the proteins from which they are derived a PET uniquely associated with a peptide fragment of said target protein that predictably results from a treatment of said biological sample.

31.   (Currently Amended) The array of claim 30, wherein said solid support comprises is beads or an array device comprising features disposed in a manner that encodes the identity of said capture agents disposed thereon.

32.   (Currently Amended) The array of claim 29, wherein said array includes 30, comprising 2 - 100 or more different capture agents.

33.   (Currently Amended) The array of claim [[29]] 30, wherein a said capture agents is a single chain antibody said array device includes a diffractive grating surface.

34.   (Currently Amended) The array of claim [[29]] 30, wherein a said capture agents are antibodies agent is an antibody or antigen binding portions thereof, and said array is an arrayed ELISA.

35-41. (Canceled)

42.   (New) A method for detecting in a biological sample the presence of plural target proteins, at least some of which comprising two or more splice variant proteins of a single DNA, the method comprising the steps of:

(1)   fragmenting proteins in the sample, using a predetermined denaturation and proteolytic protocol, to generate a solution of polypeptide analytes comprising recognition sequences displaying solvent accessible binding surfaces unambiguously indicative of the presence in the sample of the splice variant proteins from which they are derived;

(2)   providing an addressable array of plural capture agents which selectively interact with binding surfaces of said polypeptide analytes to unambiguouly indicate the presence of said splice variant proteins in said sample;

(3)   contacting said array and said solution to capture polypeptide analytes by binding interaction between respective capture agents and said binding surfaces presented by respective said analytes; and,

(4)   detecting the presence of said target splice variant proteins in the sample by detecting said binding interactions.

43.   (New) The method of claim 42, wherein the detecting step is conducted by detecting on the array the presence of captured polypeptide analytes with secondary capture agents specific for said captured polypeptide analytes and labeled with a detectable moiety.

44.   (New) The method of claim 42, wherein the addressable array of capture agents comprises plural capture agents which selectively interact with different solvent accessible binding surfaces from the same target splice variant proteins.

45.   (New) The method of claim 44, further comprising quantitating, if present, the amount of a target splice variant protein in the sample by averaging the results obtained from each said capture agent.

46.   (New) The method of claim 42, wherein said capture agents comprise antibodies.

47.   (New) The method of claim 6, wherein said subset of said capture agents have different affinity and/or avidity for said same PET.

Claims as they will appear when amended as above:

1.   **(Currently Amended)** A method for achieving high sensitivity detection and/or high accuracy quantitation of a plurality of target proteins in a biological sample, the method comprising the steps of:

   (1)   fragmenting proteins in the sample using a predetermined denaturation and proteolytic protocol to generate a solution of polypeptide analytes comprising recognition sequences comprising peptide epitope tags (PETs) displaying solvent accessible binding surfaces unambiguously indicative of the presence in the sample of the proteins from which they are derived;

   (2)   providing an addressable array of plural capture agents, which capture agents selectively interact with a peptide epitope tag (PET) of said target protein;

   (3)   contacting said array with said solution to capture polypeptide analytes by binding interaction between respective capture agents and said PETs presented by respective said analytes; and,

   (4)   detecting the presence and amounts of said target proteins in the sample by detecting said captured polypeptide analytes with secondary capture agents specific for captured polypeptide analytes and labeled with a detectable moiety.

2.   **(Currently Amended)** The method of claim 1 comprising providing an addressable array of capture agents comprising plural capture agents which selectively interact with different PETs from the same target protein and quantitating, if present, the amount of the target protein in the sample by averaging the results obtained from each said capture agent.

3.   **(Currently Amended)** The method of claim 1 wherein said capture agents comprise antibodies.

4.   **(Currently Amended)** The method of claim 1, wherein said capture agent comprise a member selected from the group consisting of non-antibody polypeptide, PNA (peptide nucleic acids), scaffolded peptide, peptidomimetic compound, polynucleotide, carbohydrates, artificial polymers, plastibody, chimeric binding agent derived from low-affinity ligand, and small organic molecules.

5.   (**Currently Amended**) The method of claim 1, wherein said secondary capture agents bind to an epitope on a said polypeptide analyte separate from said solvent accessible binding surfaces.

6.   (**Currently Amended**) The method of claim 1, wherein a subset of said capture agents bind to the same PET.

7.   (**Original**) The method of claim 1, wherein said target protein has two or more different forms within said biological sample.

8.   (**Original**) The method of claim 7, wherein said different forms include unprocessed / pro-form and processed / mature form.

9.   (**Original**) The method of claim 7, wherein said different forms include different alternative splicing forms.

10-12. (**Canceled**)

13.   (**Currently Amended**) The method of claim 7, further comprising determining the percentage of one form of said target protein as compared to the total target protein, or ratio of a first form of said target protein to a second form of said target protein.

14-15. (**Canceled**)

16.   (**Original**) The method of claim 1, wherein, for each capture agent, the method has a regression coefficient (R^2) of 0.95 or greater.

17-18. (**Canceled**)

19.   (**Original**) The method of claim 1, wherein said sample is a body fluid selected from: saliva, mucous, sweat, whole blood, serum, urine, amniotic fluid, genital fluid, fecal material, marrow, plasma, spinal fluid, pericardial fluid, gastric fluid, abdominal fluid, peritoneal fluid, pleural fluid, synovial fluid, cyst fluid, cerebrospinal fluid, lung lavage fluid, lymphatic fluid, tears, prostatitc fluid, extraction from other body parts, or secretion from other glands; or from supernatant, whole cell lysate, or cell fraction obtained by lysis and fractionation of cellular material, extract or fraction of cells obtained directly from a biological entity or cells grown in an artificial environment.

20.   (Currently Amended) The method of claim 1, wherein said sample is obtained from
       human, mouse, rat, frog, fish, fly, nematode, fission or budding yeast, or plant.

21.   (Original) The method of claim 1, wherein said sample is produced by treatment of
       membrane bound proteins.

22.   (Canceled)

23.   (Currently Amended) The method of claim 1, wherein said secondary capture agent is
       labeled with a detectable moiety selected from: an enzyme, a fluorescent label, a
       stainable dye, a chemilumninescent compound, a colloidal particle, a radioactive isotope,
       a near-infrared dye, a DNA dendrimer, a water-soluble quantum dot, a latex bead, a
       selenium particle, or a europium nanoparticle.

24.   (Currently Amended) The method of claim 23, wherein said secondary capture agent is
       labeled with a fluorophore.

25.   (Canceled)

26.   (Original) The method of claim 1, wherein said sample contains billion molar excess of
       unrelated proteins or fragments thereof relative to said target protein.

27.   (Original) The method of claim 1, wherein said PET is identified based on one or more
       of the protein sources selected from: sequenced genome or virtually translated proteome,
       virtually translated transcriptome, or mass spectrometry database of tryptic fragments.

28.   (Currently Amended) The method of claim 1, wherein one or a combination of said
       target proteins serve as a biomarker.

29.   (Canceled)

30.   (Currently Amended) An array of capture agents for detecting and quantitating plural
       target splice variant proteins within a biological sample, the array comprising a plurality
       of capture agents, each immobilized on a distinct addressable location on a solid support,
       plural of said capture agents specifically binding to recognition sequences comprising
       peptide epitope tags (PETs) displaying solvent accessible binding surfaces
       unambiguously indicative of the presence in the sample of the proteins from which they
       are derived that predictably results from a treatment of said biological sample.

31.   (Currently Amended) The array of claim 30, wherein said solid support comprises beads or an array device comprising features disposed in a manner that encodes the identity of said capture agents disposed thereon.

32.   (Currently Amended) The array of claim 30, comprising 2 - 100 or more different capture agents.

33.   (Currently Amended) The array of claim 30, wherein a said capture agents is a single chain antibody.

34.   (Currently Amended) The array of claim 30, wherein a said capture agent is an antibody or antigen binding portions thereof.

35-41. (Canceled)

42.   (New) A method for detecting in a biological sample the presence of plural target proteins, at least some of which comprising one or more splice variants of a single DNA, the method comprising the steps of:

    (1)   fragmenting proteins in the sample, using a predetermined denaturation and proteolytic protocol, to generate a solution of polypeptide analytes comprising recognition sequences displaying solvent accessible binding surfaces unambiguously indicative of the presence in the sample of splice variant proteins from which they are derived;

    (2)   providing an addressable array of plural capture agents which selectively interact with binding surfaces of said polypeptide analytes;

    (3)   contacting said array and said solution to capture polypeptide analytes by binding interaction between respective capture agents and said binding surfaces presented by respective said analytes; and,

    (4)   detecting the presence of said target splice variant proteins in the sample by detecting said binding interactions.

43.   (New) The method of claim 42, wherein the detecting step is conducted by detecting on the array the presence of captured polypeptide analytes with secondary capture agents specific for said captured polypeptide analytes and labeled with a detectable moiety.

44.   (New) The method of claim 42, wherein the addressable array of capture agents comprises plural capture agents which selectively interact with different solvent accessible binding surfaces from the same target splice variant proteins.

45.   (New) The method of claim 44, further comprising quantitating, if present, the amount of a target splice variant protein in the sample by averaging the results obtained from each said capture agent.

46.   (New) The method of claim 42, wherein said capture agents comprise antibodies.

47.   (New) The method of claim 6, wherein said subset of said capture agents have different affinity and/or avidity for said same PET.